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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant: Winn, et al.

Serial No.: 09/653,563

Filed: August 31, 2000

For: ENDOTHELIN ANTAGONISTS

Case No.: 5594.US.P9

Group Art No.: 1626

Examiner: Small, Andrea D. Souza

I hereby certify that this paper (along with any paper referred to as being attached or enclosed) is being deposited with the United States Postal Service on the date shown below with sufficient postage as first class mail in an envelope addressed to the:

Mail Stop Non-Fee Amendment  
Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450 on:

Date of Deposit: May 5, 2003

Tanya Parent

5/5/03

Date

Mail Stop Non-Fee Amendment  
Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

**TRANSMITTAL LETTER**

Dear Sir:

Enclosed herewith for the patent application identified above entitled ENDOTHELIN ANTAGONISTS are the following:

1. Preliminary Amendment (17 pages);
2. Pages 350-866 of U.S. 09/653,563; and
3. Return Receipt Postcard

The Commissioner is hereby authorized to charge any additional Filing Fees required under 37 CFR §1.16, as well as any patent application processing fees under 37 CFR §1.17 associated with this communication for which full payment had not been tendered, to Deposit Account No. 01-0025.



\* 2 3 4 9 2 \*

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Respectfully submitted,  
Winn, et al.

B. Gregory Donner  
Registration No. 34,580  
Attorney for Applicants



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**PRELIMINARY AMENDMENT**

Dear Sir:

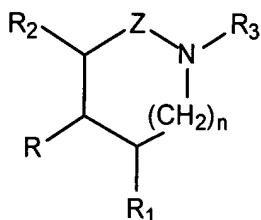
Before examination on the merits, kindly enter this amendment and reconsider this application in view of the following remarks.

**IN THE CLAIMS**

Please cancel claims 2-20 and 22-156.

Please replace claims 1, 21, and 157-160 with the correspondingly numbered claims.

1 (Amended). A compound of the formula:



wherein

Z is  $-C(R_{18})(R_{19})-$  wherein  $R_{18}$  and  $R_{19}$  are hydrogen;